# PATENT COOPERATION TREATY

| From the INTERNATIONAL SEARCHING AUTHORITY PCT   |  |  |  |  |  |  |
|--|--|--|--|--|--|--|
| To: GLAXOSMITHKLINE Corporate Intollectual Property Attn. Reed, Michael CN925.1 980 Great West Road Brentford, Middlesex TW8 9GS UNITED KINGDOM  IPM: NA CA  | DOTATION OF TRANSMITTAL OF TO THE INTERNATIONAL SEARCH REPORT AND THE WRITTEN OPINION OF THE INTERNATIONAL DEAL OF THE WRITTEN OPINION OF THE INTERNATIONAL DEAL OF THE DECLARATION APR 2005 |  |  |  |  |  |
|  | 19/04/2005   |  |  |  |  |  |
| Applicant's or agent's file reference  | FOR FURTHER ACTION See paragraphs 1 and 4 below  |  |  |  |  |  |
| MXG / PB60595 International application No.  | International filing date  |  |  |  |  |  |
| PCT/EP2004/013248  | (day/month/year) 19/11/2004  |  |  |  |  |  |
| Applicant  |  |  |  |  |  |  |
| SMITHKLINE BEECHAM CORPORATION   |  |  |  |  |  |  |
| SMITHREINE BESCHAM CORPORATION   |  |  |  |  |  |  |
| The applicant is hereby notified that the international search report and the written opinion of the International Searching Authority have been established and are transmitted herewith.  Filling of amendments and statement under Article 19: The applicant is entitled, if he so wishes, to amend the claims of the International Application (see Rule 46): When? The time limit for filing such amendments is normally 2 months from the date of transmittal of the International Search Report; however, for more details, see the notes on the accompanying sheet.  Where? Directly to the International Bureau of WIPO, 34 chemin des Colombettes  |  |  |  |  |  |  |
| 1211 Geneva 20, Switzerland, Fascimile No.: (41–22) 740.14.35  For more detailed instructions, see the notes on the accompanying sheet.  |  |  |  |  |  |  |
| <ol> <li>The applicant is hereby notified that no international search report will be established and that the declaration under Article 17(2)(a) to that effect and the written opinion of the international Searching Authority are transmitted herewith.</li> <li>With regard to the protest against payment of (an) additional fee(s) under Rule 40.2, the applicant is notified that:         <ul> <li>the protest together with the decision thereon has been transmitted to the International Bureau together with the applicant's request to forward the texts of both the protest and the decision thereon to the designated Offices.</li> <li>no decision has been made yet on the protest; the applicant will be notified as soon as a decision is made.</li> </ul> </li> </ol> |  |  |  |  |  |  |
| 4. Reminders   |  |  |  |  |  |  |
| Shortly after the expiration of <b>18</b> months from the priority date, the international application will be published by the International Bureau. If the applicant wishes to avoid or postpone publication, a notice of withdrawal of the international application, or of the priority claim, must reach the international Bureau as provided in Rules 90 <i>bis.</i> 1 and 90 <i>bis.</i> 3, respectively, before the completion of the technical preparations for international publication.  The applicant may submit comments on an informal basis on the written opinion of the international Searching Authority to the   |  |  |  |  |  |  |
| international Bureau. The International Bureau will send a copy of<br>International preliminary examination report has been or is to be e<br>the public but not before the expiration of 30 months from the prior  | such comments to all designated Offices unless an<br>stablished. These comments would also be made available to  |  |  |  |  |  |
| Within 19 months from the priority date, but only in respect of sor examination must be filed if the applicant wishes to postpone the date (in some Offices even later); otherwise, the applicant must, wacts for entry into the national phase before those designated Offi   | entry into the national phase until 30 months, from the priority<br>within 20 months from the priority date, perform the prescribed  |  |  |  |  |  |
| In respect of other designated Offices, the time fimit of 30 months months.  | (or later) will apply even if no demand is filed within 19   |  |  |  |  |  |
| See the Annex to Form PCT/IE/301 and, for details about the app Guide, Volume II, National Chapters and the WIPO Internet site.  | licable time limits, Office by Office, see the PCT Applicant's   |  |  |  |  |  |
| Name and malling address of the International Searching Authority  | Authorized officer   |  |  |  |  |  |
| European Patent Office, P.B. 5818 Patentlaan 2<br>NL-2280 HV Rijswijk<br>Tel. (+31~70) 340-2040, Tx. 31 651 epo nl,<br>Fax: (+31-70) 340-3016  | Joëlle Gerber  |  |  |  |  |  |

# PATENT COOPERATION TREATY

# **PCT**

## INTERNATIONAL SEARCH REPORT

(PCT Article 18 and Rules 43 and 44)

| Applicant's or agent's file reference  | FOR FURTHER   |               | see Form PCT/ISA/220   |
|--|---|---------------|--|
| MXG/PB60595  | ACTION  | as well a     | as, where applicable, Item 5 below.  |
| International application No.  | International filing date (day/month  | /year)        | (Earliest) Priority Date (day/month/year)  |
| PCT/EP2004/013248  | 19/11/2004  |               | 21/11/2003   |
| Applicant  |   |               |  |
| SMITHKLINE BEBCHAM CORPORA   | ATION   | ii            |  |
| This international Search Report has been according to Article 18. A copy is being tra | n prepared by this international Sear<br>ansmitted to the international Bureau              | ching Autho   | prity and is transmitted to the applicant  |
| This International Search Report consists  | of a total of she   | ets.          |  |
| X It is also accompanied by  | a copy of each prior art document d   | ted in this r | report.  |
| 1. Basis of the report   |   |               |  |
|  | International search was carried out<br>ess otherwise indicated under this its              |               | is of the international application in the   |
| The international this Authority (Ru   |   | of a transla  | tion of the international application furnished to   |
| b. With regard to any nucleo   | otide and/or amino acid sequence  | disclosed b   | n the International application, see Box No. I.  |
| 2. K Certain claims were fou   | nd unsearchable (See Box II).   |               | 1 9 APR 2005   |
| 3. Unity of invention is lac   | king (see Box III).   |               | ELESALIS NISP  |
| 4. With regard to the title,   |   |               | 1  |
| the text is approved as su   | •   |               |  |
| $oxed{X}$ the text has been established USE OF (2S, 3S) 2-(3-0)                        | thed by this Authority to read as follo   |               | 2 Mapped That  |
| 006 OF (28, 35) 2-(3-(   | WHOKORWEWIP) = 3, 3, 3-1K   | TMÜTUTT       | 7-5-MOKAUOTIMOT  |
|  |   |               |  |
|  |   |               |  |
|  |   |               |  |
|  |   |               |  |
|  |   |               |  |
| 5. With regard to the abstract,  |   |               |  |
| X the text is approved as su   | • • •   |               |  |
|  |   |               | y as it appears in Box No. IV. The applicant<br>h report, submit comments to this Authority. |
| 6. With regard to the drawings,  |   |               |  |
| a. the figure of the drawings to be p  | sublished with the abstract is Figure (   | <b>V</b> o    | <u></u>  |
| as suggested by  | the applicant.  |               |  |
|  | s Authority, because the applicant fa   |               | •  |
| l . — — ·  | <ul> <li>Authority, because this figure better<br/>e published with the abstract</li> </ul> | rcharacter    | izes me invention.   |
| b. Indite of the lightes is to b   | e published with the abstract.  |               |  |

international Application No PCT/EP2004/013248

A. CLASSIFICATION OF SUBJECT MATTER IPC 7 A61K31/5375 A61P25/14

According to International Patent Classification (IPC) onto both national classification and IPC

B. FIELDS SEARCHED

Minimum documentation searched (classification system followed by classification symbols) IPC 7 A61K

Documentation searched other than minimum documentation to the extent that such documents are included in the fields searched

Electronic data base consulted during the international search (name of data base and, where practical, search terms used)

EPO-Internal, CHEM ABS Data, EMBASE, MEDLINE, SCISEARCH, BIOSIS

| Calegory * | Citation of document, with indication, where appropriate, of the relevant passages   | Aelevant to claim No. |  |  |
|------------|--|-----------------------|--|--|
| A          | US 2003/064988 A1 (MORGAN PHILLIP<br>FREDERICK ET AL) 3 April 2003 (2003-04-03)<br>paragraph '0006! - paragraph '0010!<br>paragraph '0023! |                       |  |  |
| A          | US 6 337 328 B1 (FANG QUN KEVIN ET AL)<br>8 January 2002 (2002-01-08)<br>column 4, line 40 - line 63                                       |                       |  |  |
| A          | WO 01/62257 A (SEPRACOR INC) 30 August 2001 (2001-08-30) page 4, line 29 - page 5, line 1 page 7, line 26 - line 31                        |                       |  |  |
|            | -/ <b></b>   |                       |  |  |
|            |  |                       |  |  |
|            |  |                       |  |  |

| X Further documents are listed in the continuation of box C   | Palent family members are listed in annex.   |
|---|--|
| <ul> <li>Special categories of cited documents:</li> <li>"A" document defining the general state of the last which is not considered to be of particular relevance</li> <li>"E" earlier document but published on or after the international filling date</li> <li>"L" document which may throw doubts on priority claim(s) or which is cited to establish the publication date of another citation or other special reason (as specified)</li> <li>"O" document reterring to an oral discheure, use, exhibition or other means</li> <li>"D" document published prior to the international filling date but later than the priority date claimed</li> </ul> | <ul> <li>'T' later document published after the international filing date or priority date and not in conflict with the application but cited to understand the principle or theory underlying the invention.</li> <li>'X' document of particular relevance; the claimed invention cannot be considered novel or cannot be considered to involve an inventive step when the document is taken alone.</li> <li>'Y' document of particular relevance; the claimed invention cannot be considered to involve an inventive step when the document is combined with one or more other such document is combined with one or more other such documents, such combination being obvious to a person skilled in the art.</li> <li>'&amp;' document member of the same patent family</li> </ul> |
| Date of the actual completion of the international search   | Date of mailing of the international search report   |
| 13 Apr11 2005   | 19/04/2005   |
| Name and mailing address of the ISA<br>European Patent Office, P.B. 5616 Patentlaah 2<br>NL - 2280 HV Rijswijk  | Authorized officer   |
| Tol (+31-70) 340-2040. Tx. 31 651 epo nl.<br>Fax: (+31-70) 340-3016   | Bonzano, C   |

International Application No PCT/EP2004/012249

|   |   | PCT/EP2004/013248     |  |  |  |  |
|---|---|-----------------------|--|--|--|--|
| Continuation) DOCUMENTS CONSIDERED TO BE RELEVANT |   |                       |  |  |  |  |
| ategory *   | Citation of document, with indication, where appropriate, of the relevant passages  | Relevant to claim No. |  |  |  |  |
| A   | WO 99/37305 A (GLAXO GROUP LIMITED;<br>MORGAN, PHILLIP, FREDERICK; MUSSO, DAVID,<br>LEE; PA) 29 July 1999 (1999-07-29)<br>cited in the application<br>abstract<br>page 2, line 5 - line 14  |                       |  |  |  |  |
| Y /   | KELLEY JAMES L ET AL: "-2-(3,5-difluorophenyl)-3,5-dimethyl-2-morpholinol: A novel antidepressant agent and selective inhibitor of norepinephrine uptake" JOURNAL OF MEDICINAL CHEMISTRY, AMERICAN CHEMICAL SOCIETY. WASHINGTON, US, vol. 39, no. 2, 1996, pages 347-349, XP002164358 ISSN: 0022-2623 page 347, column 2, paragraph 3 | 1-12                  |  |  |  |  |
| Y   | EP 1 336 406 A (SOLVAY PHARM BV 'NL!) 20 August 2003 (2003-08-20) paragraph '0013! paragraph '0001!   | 1-12                  |  |  |  |  |
|   |   |                       |  |  |  |  |

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MAY-17-2006 12:06 From:GSK

From PCT/ISA/210 (continuation of second sheet) (January 2004)

International application No. PCT/EP2004/013248

| Box II Observations where certain claims were found unsearchable (Continuation of Item 2 of first sheet)  |
|---|
| This international Search Report has not been established in respect of certain claims under Article 17(2)(a) for the following reasons:  |
| 1. X Claims Nos.: because they relate to subject matter not required to be searched by this Authority, namely:  |
| Although claims 10-12 are directed to a method of treatment of the human/animal body, the search has been carried out and based on the alleged effects of the compound/composition.   |
| 2. Ctalms Nos.: because they relate to parts of the International Application that do not comply with the prescribed requirements to such an extent that no meaningful international Search can be carried out, specifically: |
|   |
| 3. Claims Nos.:  because they are dependent claims and are not drafted in accordance with the second and third sentences of Rule 6.4(a).  |
| Box III Observations where unity of invention is lacking (Continuation of Item 3 of first sheet)  |
| This International Searching Authority found multiple Inventions in this International application, as follows:   |
|   |
| As all required additional search fees were timely paid by the applicant, this international Search Report covers all searchable claims.  |
| 2. As all searchable claims could be searched without effort justifying an additional fee, this Authority did not invite payment of any additional fee.   |
| 3. As only some of the required additional search fees were timely paid by the applicant, this International Search Report covers only those claims for which fees were paid, specifically claims Nos.:                       |
| 4. No required additional search fees were timely paid by the applicant. Consequently, this international Search Report is restricted to the invention first mentioned in the claims; it is covered by claims Nos.:           |
| Remark on Protest  The additional search fees were accompanied by the applicant's protest.  No protest accompanied the payment of additional search fees.   |

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## INTERNATIONAL SEARCH REPORT

919 483 7977

Information on patent family members

International Application No PCT/EP2004/013248

|      |                                  |    | ·                   |  |   | PC 1/EPZ   | 004/013248   |
|------|----------------------------------|----|---------------------|--|---|--|--|
|      | ent document<br>in search report |    | Publication<br>date |  | Patent family<br>member(s)  |  | Publication<br>date  |
| US 1 | 2003064988                       | A1 | 03-04-2003          | US<br>US<br>US<br>US<br>US   | 2002019396<br>6274579<br>2003027827<br>2003083330<br>2003032643   | ) B1<br>/ A1<br>) A1   | 14-02-2002<br>14-08-2001<br>06-02-2003<br>01-05-2003<br>13-02-2003   |
| US ( | 6337328                          | B1 | 08-01-2002          | US<br>AU<br>CA<br>CZ<br>EP<br>HU<br>JP<br>PL<br>WO<br>CA<br>EP<br>JP<br>AU<br>WO | 6342496<br>6926806<br>2400482<br>20022857<br>1259243<br>0300036<br>2003529563<br>357389<br>0162257<br>2002052341<br>2362361<br>1165086<br>2004513063<br>2002052346<br>775642<br>3505506 | ) A<br>2 A1<br>7 A3<br>3 A2<br>3 T<br>9 A1<br>7 A2<br>1 A1<br>1 A1<br>5 A2<br>1 T<br>2 B2            | 29-01-2002<br>03-09-2001<br>30-08-2001<br>14-01-2004<br>27-11-2002<br>28-05-2003<br>07-10-2003<br>26+07-2004<br>30+08-2001<br>02-05-2002<br>08-09-2000<br>02-01-2002<br>30-04-2004<br>02-05-2002<br>12-08-2004<br>21-09-2000<br>08-09-2000 |
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information on patent family members

International Application No
PCT/EP2004/013248

| Patent document<br>died in search report |   | Publication<br>date |  | Patent (amily<br>member(s)   |                           | Publication date   |
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| WO 9937305                               | A |                     | PL<br>SK<br>TR<br>WO<br>US<br>US       | 342012<br>10912000<br>200002126<br>9937305<br>6274579<br>2002019396              | A3<br>T2<br>A1<br>B1      | 07-05-2001<br>09-04-2001<br>21-12-2000<br>29-07-1999<br>14-08-2001<br>14-02-2002               |
| EP 1336406                               | A | 20-08-2003          | EP<br>AU<br>BR<br>CA<br>WO<br>EP<br>US | 1336406<br>2003208714<br>0306147<br>2462688<br>03068207<br>1476148<br>2005038015 | A1<br>A<br>A1<br>A2<br>A2 | 20-08-2003<br>04-09-2003<br>19-10-2004<br>21-08-2003<br>21-08-2003<br>17-11-2004<br>17-02-2005 |